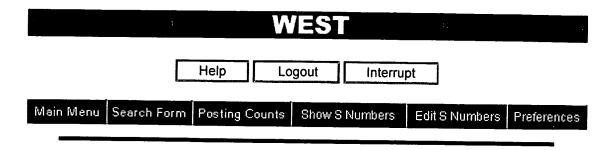
DB Name	<u>Query</u>	Hit Count	Set Name
JPAB,EPAB,DWPI	'alpha4beta7' same (antibod\$ or hybridoma\$)	2	<u>L3</u>
JPAB,EPAB,DWPI	'act-1' same (antibod\$ or hybridoma\$)	2	<u>L2</u>
USPT	'act-1' same (antibod\$ or hybridoma\$)	1	<u>L1</u>



Search Results -

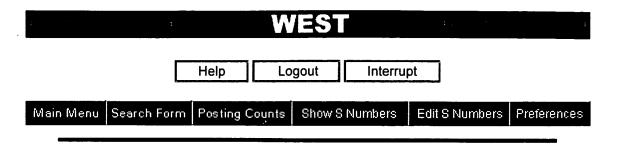
Term	Documents
ACT-1.DWPI,EPAB,JPAB.	4
ACT-1S	0
ANTIBOD\$	0
ANTIBOD.DWPI,EPAB,JPAB.	49
ANTIBODANTIBODA.DWPI,EPAB,JPAB.	1
ANTIBODES.DWPI,EPAB,JPAB.	11
ANTIBODI.DWPI,EPAB,JPAB.	10
ANTIBODIC.DWPI,EPAB,JPAB.	1
ANTIBODIDES.DWPI,EPAB,JPAB.	
ANTIBODIE.DWPI,EPAB,JPAB.	7
('ACT-1' SAME (ANTIBOD\$ OR HYBRIDOMA\$)).JPAB,EPAB,DWPI.	2

There are more results than shown above. Click here to view the entire set.

Database:	US Patents Full-Text Database US Pre-Grant Publication Full-Text Database JPO Abstracts Database EPO Abstracts Database Derwent World Patents Index IBM Technical Disclosure Bulletins
Refine S	'act-1' same (antibod\$ or hybridoma\$) earch: Clear
	Search History

Today's Date: 4/15/2001

DB Name	Query	Hit Count	Set Name
JPAB,EPAB,DWPI	'act-1' same (antibod\$ or hybridoma\$)	2	<u>L2</u>
USPT	'act-1' same (antibod\$ or hybridoma\$)	1	<u>L1</u>



Search Results -

Term	Documents
ALPHA4BETA7.DWPI,EPAB,JPAB.	2
ALPHA4BETA7S	0
ANTIBOD\$	0
ANTIBOD.DWPI,EPAB,JPAB.	49
ANTIBODANTIBODA.DWPI,EPAB,JPAB.	1
ANTIBODES.DWPI,EPAB,JPAB.	11
ANTIBODI.DWPI,EPAB,JPAB.	10
ANTIBODIC.DWPI,EPAB,JPAB.	1
ANTIBODIDES.DWPI,EPAB,JPAB.	1
ANTIBODIE.DWPI,EPAB,JPAB.	7
('ALPHA4BETA7' SAME (ANTIBOD\$ OR HYBRIDOMA\$)).JPAB,EPAB,DWPI.	2

There are more results than shown above. Click here to view the entire set.

Refine S	earcn.			Clear	and consists on the last and another to the
Pofino S	oorob:	'alpha4beta7' same (antib hybridoma\$)	od\$ or	Class	
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	EPO Abs	stracts Database			
		Grant Publication Full-Text Database tracts Database	3		
		nts Full-Text Database			

Today's Date: 4/15/2001

Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	ROME	Draw Desc	Image
									_	0.01 0.030	1111985

Generate Collection

Term	Documents
ACT-1.DWPI,EPAB,JPAB.	4
ACT-1S	0
ANTIBOD\$	0
ANTIBOD.DWPI,EPAB,JPAB.	49
ANTIBODANTIBODA.DWPI,EPAB,JPAB.	1
ANTIBODES.DWPI,EPAB,JPAB.	11
ANTIBODI.DWPI,EPAB,JPAB.	10
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ANTIBODIDES.DWPI,EPAB,JPAB.	1
ANTIBODIE.DWPI,EPAB,JPAB.	7
('ACT-1' SAME (ANTIBOD\$ OR HYBRIDOMA\$)).JPAB,EPAB,DWPI.	2

There are more results than shown above. Click here to view the entire set.

Display	10 Documents, starting with Document:	2
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Display Format: CIT Change Format

WEST

End of Result Set

Generate Collection

L1: Entry 1 of 1

File: USPT

Jul 13, 1999

DOCUMENT-IDENTIFIER: US 5922676 A

TITLE: Methods of inhibiting cancer by using superfibronectin

DEPR:

To provide a basis for the in vitro studies of FIG. 7, the integrin surface expression profile on the tumor cell lines was determined by flow cytometric analysis as described in Pasqualini, R., et al., J. Cell. Biol. 125:447-460 (1994). Monoclonal antibodies against the human .alpha..sub.2 (AK-7), and .alpha..sub.4 (9F10) subunits were from Pharmingen (La Jolla, Calif.), anti-.beta..sub.3 (CD-61) was from Becton and Dickinson (San Jose, Calif.), anti-.beta..sub.4 (3E1) was from Gibco BRL (Bethesda, Md.), anti-.alpha..sub.5 from Oncogene Science (Cambridge, Mass.), anti-.alpha..sub.V from Chemicon (Temecula, Calif.), anti-.beta..sub.1 (TS2/16), anti-.beta..sub.5 (IA9), anti-.alpha..sub.1 (TS2/7), anti-.alpha..sub.6 (BQ16), and anti-.alpha..sub.3 (IVA5) were a gift from Dr. Martin Hemler (Dana Farber Cancer Institute, Harvard Medical School); the anti-.beta..sub.8 (SN-1) was from Dr. Robert Pytela (University of California, San Francisco), anti-.beta..sub.6 (9G6) and anti .alpha..sub.9 (Y9A2) were from Dr. Dean Sheppard (University of California, San Francisco), and the anti-.alpha..sub.4 .beta..sub.7 (Act-1) antibody was from Dr. Andrew Lazarovits (University of Western Ontario, Canada).

WEST

Generate Collection

Search Results - Record(s) 1 through 2 of 2 returned.

☐ 1. Document ID: WO 9806248 A2

L2: Entry 1 of 2

File: EPAB

Feb 19, 1998

PUB-NO: WO009806248A2

DOCUMENT-IDENTIFIER: WO 9806248 A2 TITLE: TITLE DATA NOT AVAILABLE

PUBN-DATE: February 19, 1998

Full Title Citation Front Review Classification Date Reference Claims KMC Draw Desc Image	Full	Title	Citation	Front	Review	Classification	Date	Reference	Claims	KWC	Draw, Desc	Image
---	------	-------	----------	-------	--------	----------------	------	-----------	--------	-----	------------	-------

Document ID: AU 730326 B, WO 9806248 A2, AU 9739728 A, EP 918797 A2, CN 1227607 A, BR 9711079 A, MX 9901462 A1, NZ 334226 A

L2: Entry 2 of 2

File: DWPI

Mar 1, 2001

DERWENT-ACC-NO: 1998-159172

DERWENT-WEEK: 200117

COPYRIGHT 2001 DERWENT INFORMATION LTD

TITLE: Humanised immunoglobulin reactive with alpha-4-beta-7 integrin - used for treating inflammatory disease, pancreatitis, diabetes, asthma, graft versus host disease and sarcoidosis

INVENTOR: BENDIG, M M; JONES, S T ; NEWMAN, W ; PONATH, P D ; RINGLER, D J ; SALDANHA, J ; TARRAN JONES, S

PRIORITY-DATA: 1996US-0700737 (August 15, 1996)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
AU 730326 B	March 1, 2001	N/A	000	A61K039/00
WO 9806248 A2	February 19, 1998	E	145	C12N000/00
AU 9739728 A	March 6, 1998	N/A	000	A61K039/00
EP 918797 A2	June 2, 1999	E	000	C07K016/28
CN 1227607 A	September 1, 1999 '	N/A	000	C12N015/62
BR 9711079 A	June 6, 2000	N/A	000	C07K016/46
MX 9901462 A1	May 1, 1999	N/A	000	A61K045/00
NZ 334226 A	October 27, 2000	N/A	000	C12N005/10

INT-CL (IPC): A61K 39/00; A61K 39/395; A61K 45/00; C07K 14/705; C07K 14/78; C07K 16/28; C07K 16/46; C12N 0/00; C12N 5/10; C12N 5/18; C12N 5/22; C12N 15/13; C12N 15/62; C12N 15/85; G01N 33/577

WEST

Your wildcard search against 2000 terms has yielded the results below

Search for additional matches among the next 2000 terms

Generate Collection

Search Results - Record(s) 1 through 1 of 1 returned.

☐ 1. Document ID: US 5922676 A

L1: Entry 1 of 1

File: USPT

Jul 13, 1999

US-PAT-NO: 5922676

DOCUMENT-IDENTIFIER: US 5922676 A

TITLE: Methods of inhibiting cancer by using superfibronectin

DATE-ISSUED: July 13, 1999

INVENTOR-INFORMATION:

NAME

CITY

STATE ZIP CODE

COUNTRY

Pasqualini; Renata

Solana Beach

CA N/A

N/A

Ruoslahti; Erkki

Rancho Santa Fe

CA N/A

N/A

US-CL-CURRENT: 514/12; 424/499, 435/402, 514/2, 530/324

Full Title Citation Front Review Classification Date Reference Claims KWC Draw Desc Image

Generate Collection

Term	Documents
ACT-1.USPT.	15
ACT-1S]0
ANTIBOD\$	0
ANTIBOD.USPT.	223
ANTIBODAY.USPT.]1
ANTIBODEES.USPT.	1
ANTIBODEIS.USPT.	1
ANTIBODES.USPT.	44
ANTIBODEY.USPT.	1
ANTIBODFIES.USPT.	1
('ACT-1' SAME (ANTIBOD\$ OR HYBRIDOMA\$)).USPT.	1

There are more results than shown above. Click here to view the entire set.

Display 10 Documents, starting with Document: 1

Display Format: CIT Change Format

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>>>'IALOG' not recognized as set or accession number
? set hi ;set hi
       15apr01 11:37:04 User208760 Session D1827.1
            $0.41 0.116 DialUnits File1
     $0.41 Estimated cost File1
     $0.05 TYMNET
     $0.46 Estimated cost this search
     $0.46 Estimated total session cost 0.116 DialUnits
File 410:Chronolog(R) 1981-2001 Mar/Apr
       (c) 2001 The Dialog Corporation
      Set Items Description
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     $0.01 TYMNET
     $0.01 Estimated cost this search
     $0.47 Estimated total session cost 0.172 DialUnits
SYSTEM:OS - DIALOG OneSearch
 File 5:Biosis Previews(R) 1969-2001/Apr W2
         (c) 2001 BIOSIS
  File 73:EMBASE 1974-2001/Apr W2
         (c) 2001 Elsevier Science B.V.
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see Help News73.
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         (c) 2001 AMERICAN CHEMICAL SOCIETY
*File 399: Use is subject to the terms of your user/customer agreement.
RANK charge added; see HELP RATES 399.
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         198801 ACT
        7661201 1
197 ACT(W)1
     S1
? s act(w)1 (20n)(antibod$ or hybridoma?)
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        7661201 1
              0 ANTIBOD$
          44215 HYBRIDOMA?
     S2
              0 ACT(W)1 (20N) (ANTIBOD$ OR HYBRIDOMA?)
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? s act(w)1 and (antibod\$ or hybridoma? or adhesion or integrin?)

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           44215 HYBRIDOMA?
          290335 ADHESION
           56376 INTEGRIN?
      S3
              20 ACT(W)1 AND (ANTIBOD$ OR HYBRIDOMA? OR ADHESION OR
? rd s3
...completed examining records
              15 RD S3 (unique items)
? t s4/3/all
 4/3/1
           (Item 1 from file: 5)
DIALOG(R) File
                5:Biosis Previews(R)
(c) 2001 BIOSIS. All rts. reserv.
          BIOSIS NO.: 199699247431
Distribution of beta-7 integrins in human intestinal mucosa and
  organized gut-associated lymphoid tissue.
AUTHOR: Farstad I N(a); Halstensen T H; Lien B; Kilshaw P J; Lazarovitz A I
  ; Brandtzaeg P
AUTHOR ADDRESS: (a) Inst. Pathol., The Natl. Hosp., Rikshospitalet, N-0027
  Oslo**Norway
JOURNAL: Immunology 89 (2):p227-237 1996
ISSN: 0019-2805
DOCUMENT TYPE: Article
RECORD TYPE: Abstract
LANGUAGE: English
           (Item 2 from file: 5)
DIALOG(R) File 5: Biosis Previews(R)
(c) 2001 BIOSIS. All rts. reserv.
          BIOSIS NO.: 199699200403
Integrin alpha-4-beta-7 mediates human eosinophil interaction with
  MAdCAM-1, VCAM-1 and fibronectin.
AUTHOR: Walsh G M(a); Symon F A; Lazarovits A I; Wardlaw A J
AUTHOR ADDRESS: (a) Dep. Respiratory Med., Univ. Leicester Med. Sch.,
  Glenfield Hospital, Leicester LE3 9QP**UK
JOURNAL: Immunology 89 (1):p112-119 1996
ISSN: 0019-2805
DOCUMENT TYPE: Article
RECORD TYPE: Abstract.
LANGUAGE: English
 4/3/3
           (Item 3 from file: 5)
DIALOG(R) File
               5:Biosis Previews(R)
(c) 2001 BIOSIS. All rts. reserv.
10579256
          BIOSIS NO.: 199699200401
Expression and function of alpha-4/beta-7 integrin on human natural
  killer cells.
AUTHOR: Perez-Villar J J(a); Zapata J M; Melero I; Postigo A;
  Sanchez-Madrid F; Lopez-Botet M
AUTHOR ADDRESS: (a) Serv. Immunologia, Hospital de la Princesa, Diego de
  Leon 62, 28006 Madrid**Spain
JOURNAL: Immunology 89 (1):p96-104 1996
ISSN: 0019-2805
DOCUMENT TYPE: Article
```

RECORD TYPE: Abstract LANGUAGE: English (Item 4 from file: 5) DIALOG(R) File 5:Biosis Previews(R) (c) 2001 BIOSIS. All rts. reserv. BIOSIS NO.: 199699103622 Specific inhibition of T lymphocyte coactivation by triggering integrin beta-1 reveals convergence of beta-1, beta-2, and beta-7 signaling pathways. AUTHOR: Woodside Darren G; Teague T Kent; McIntyre Bradley W(a) AUTHOR ADDRESS: (a) Univ. Texas, M.D. Anderson Cancer Cent., Dep. Immunology, Box 180, 1515 Holcombe Boulevard, Hous**USA JOURNAL: Journal of Immunology 157 (2):p700-706 1996 ISSN: 0022-1767 DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English (Item 5 from file: 5) 5:Biosis Previews(R) DIALOG(R) File (c) 2001 BIOSIS. All rts. reserv. BIOSIS NO.: 199497074987 Differential expression in rheumatoid synovium and synovial fluid of alpha-4-beta-7 integrin. A novel receptor for Fibronectin and vascular cell adhesion molecule-1. AUTHOR: Lazarovits Andrew I(a); Karsh Jacob AUTHOR ADDRESS: (a) Univ. Hosp., Room 4TU46, Box 5339, 339 Windermere Rd., London, ON N6A 5A5**Canada JOURNAL: Journal of Immunology 151 (11):p6482-6489 1993 ISSN: 0022-1767 DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English 4/3/6 (Item 6 from file: 5) DIALOG(R) File 5: Biosis Previews(R) (c) 2001 BIOSIS. All rts. reserv. 08920904 BIOSIS NO.: 199396072405 CD53, a protein with four membrane-spanning domains, mediates signal transduction in human monocytes and B cells. AUTHOR: Olweus Johanna(a); Lund-Johansen Fridtjof; Horejsi Vaclav AUTHOR ADDRESS: (a) Res. Dep., Becton Dickinson Immunocytometry Systems, 2350 Qume Drive, San Jose, CA 95131-1807**USA JOURNAL: Journal of Immunology 151 (2):p707-716 1993 ISSN: 0022-1767 DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English (Item 7 from file: 5) DIALOG(R) File 5: Biosis Previews(R) (c) 2001 BIOSIS. All rts. reserv. BIOSIS NO.: 199396072404 Selective expression of integrin alpha-4-beta-7 on a subset of human CD4 positive memory T cells with hallmarks of gut-trophism.

AUTHOR: Schweighoffer Tamas(a); Tanaka Yoshiya; Tidswell Mark; Erle David J; Horgan Kevin J; Luce Gale E Ginther; Lazarovits Andrew I; Buck David;

```
Shaw Stephen
 AUTHOR ADDRESS: (a) Exp. Immunol. Branch, NCI, NIH, Build. 10, Room 4B17,
   Bethesda, MD 20892**USA
 JOURNAL: Journal of Immunology 151 (2):p717-729 1993
 ISSN: 0022-1767
 DOCUMENT TYPE: Article
 RECORD TYPE: Abstract
 LANGUAGE: English
            (Item 1 from file: 73)
  4/3/8
 DIALOG(R) File 73: EMBASE
 (c) 2001 Elsevier Science B.V. All rts. reserv.
 06647794
              EMBASE No: 1996312652
   Distribution of betainf 7 integrins in human intestinal mucosa and
 organized gut-associated lymphoid tissue
   Farstad I.N.; Halstensen T.S.; Lien B.; Kilshaw P.J.; Lazarovitz A.I.;
 Brandtzaeg P.
   Institute of Pathology, National Hospital, Rikshospitalet, N-0027 Oslo
   Immunology (IMMUNOLOGY) (United Kingdom) 1996, 89/2 (227-237)
  CODEN: IMMUA
                  ISSN: 0019-2805
  DOCUMENT TYPE: Journal; Article
  LANGUAGE: ENGLISH
                      SUMMARY LANGUAGE: ENGLISH
  4/3/9
            (Item 2 from file: 73)
 DIALOG(R) File 73: EMBASE
 (c) 2001 Elsevier Science B.V. All rts. reserv.
             EMBASE No: 1996262400
  Integrin alphainf 4betainf 7 mediates human eosinophil interaction
with MAdCAM-1, VCAM-1 and fibronectin
  Walsh G.M.; Symon F.A.; Lazarovits A.I.; Wardlaw A.J.
  Department of Respiratory Medicine, University Leicester Medical School,
  Glenfield Hospital, Leicester LE3 9QP United Kingdom
  Immunology (IMMUNOLOGY) (United Kingdom) 1996, 89/1 (112-119)
  CODEN: IMMUA
                 ISSN: 0019-2805
  DOCUMENT TYPE: Journal; Article
  LANGUAGE: ENGLISH
                     SUMMARY LANGUAGE: ENGLISH
 4/3/10
            (Item 3 from file: 73)
DIALOG(R) File 73: EMBASE
(c) 2001 Elsevier Science B.V. All rts. reserv.
             EMBASE No: 1996262398
  Expression and function of alphainf 4/betainf 7 integrin on human
natural killer cells
  Perez-Villar J.J.; Zapata J.M.; Melero I.; Postigo A.; Sanchez-Madrid F.;
Lopez-Botet M.
  Servicio de Inmunologia, Hospital de la Princesa, Diego de Leon 62,28006
  Madrid Spain
  Immunology ( IMMUNOLOGY ) (United Kingdom) 1996, 89/1 (96-104)
  CODEN: IMMUA
                 ISSN: 0019-2805
  DOCUMENT TYPE: Journal; Article
  LANGUAGE: ENGLISH
                     SUMMARY LANGUAGE: ENGLISH
 4/3/11
            (Item 4 from file: 73)
DIALOG(R) File 73: EMBASE
(c) 2001 Elsevier Science B.V. All rts. reserv.
06549143
             EMBASE No: 1996211077
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Specific inhibition of T lymphocyte coactivation by triggering

integrin betainf 1 reveals convergence of betainf 1, betainf 2, and betainf 7 signaling pathways Woodside D.G.; Teague T.K.; McIntyre B.W. Department of Immunology, M. D. Andreson Cancer Center, University of Texas, 1515 Holcombe Boulevard, Houston, TX 77030 United States Journal of Immunology (J. IMMUNOL.) (United States) 1996, 157/2 (700 - 706)CODEN: JOIMA ISSN: 0022-1767 DOCUMENT TYPE: Journal; Article LANGUAGE: ENGLISH SUMMARY LANGUAGE: ENGLISH 4/3/12 (Item 5 from file: 73) DIALOG(R) File 73: EMBASE (c) 2001 Elsevier Science B.V. All rts. reserv. 05447652 EMBASE No: 1993215751 Selective expression of integrin alpha4beta7 on a subset of human CD4sup + memory T cells with hallmarks of gut-trophism Schweighoffer T.; Tanaka Y.; Tidswell M.; Erle D.J.; Horgan K.J.; Luce G.E.G.; Lazarovits A.I.; Buck D.; Shaw S.

Experimental Immunology Branch, NCI, NIH, Bethesda, MD 20892 United States

Journal of Immunology (J. IMMUNOL.) (United States) 1993, 151/2 (717 - 729)

CODEN: JOIMA ISSN: 0022-1767 DOCUMENT TYPE: Journal; Article

LANGUAGE: ENGLISH SUMMARY LANGUAGE: ENGLISH

4/3/13 (Item 1 from file: 155) DIALOG(R) File 155: MEDLINE(R) (c) format only 2000 Dialog Corporation. All rts. reserv.

09195954 97376878

Structure-function analysis of the integrin beta 7 subunit: identification of domains involved in adhesion to MAdCAM-1.

Tidswell M; Pachynski R; Wu SW; Qiu SQ; Dunham E; Cochran N; Briskin MJ; Kilshaw PJ; Lazarovits AI; Andrew DP; Butcher EC; Yednock TA; Erle DJ Lung Biology Center, Department of Medicine, University of California, San Francisco 94143, USA. easd@itsa.ucsf.edu

Journal of immunology (UNITED STATES) Aug 1 1997, 159 (3) p1497-505, ISSN 0022-1767. Journal Code: IFB

Contract/Grant No.: K08HL03230, HL, NHLBI; R01HL52004, HL, NHLBI; AI37832 , AI, NIAID

Languages: ENGLISH

Document type: JOURNAL ARTICLE

(Item 2 from file: 155) DIALOG(R) File 155: MEDLINE(R) (c) format only 2000 Dialog Corporation. All rts. reserv.

08147514 95261715

Integrin alpha 4 beta 7 co-stimulation of human peripheral blood T cell proliferation.

Teague TK; Lazarovits AI; McIntyre BW

Department of Immunology, University of Texas M. D. Anderson Cancer, Center, Houston 77030, USA.

Cell adhesion and communication (SWITZERLAND) Dec 1994, 2 (6) p539-47, ISSN 1061-5385 Journal Code: B4A

Contract/Grant No.: CA62596, CA, NCI

Languages: ENGLISH

Document type: JOURNAL ARTICLE

4/3/15 (Item 3 from file: 155)
DIALOG(R)File 155:MEDLINE(R)
(c) format only 2000 Dialog Corporation. All rts. reserv.

05324396 88236439

Human cytotrophoblastic antigens defined by monoclonal antibodies.

Abe Y; Okamura K; Hamazaki Y; Wada Y; Yajima A

Department of Obstetrics and Gynecology, Tohoku University School of Medicine, Sendai.

Tohoku journal of experimental medicine (JAPAN) Mar 1988, 154 (3) p245-51, ISSN 0040-8727 Journal Code: VTF

Languages: ENGLISH

Document type: JOURNAL ARTICLE

4/7/4 (Item 4 from file: 5)
DIALOG(R)File 5:Biosis Previews(R)
(c) 2001 BIOSIS. All rts. reserv.

10482477 BIOSIS NO.: 199699103622

Specific inhibition of T lymphocyte coactivation by triggering integrin beta-1 reveals convergence of beta-1, beta-2, and beta-7 signaling pathways.

AUTHOR: Woodside Darren G; Teague T Kent; McIntyre Bradley W(a) AUTHOR ADDRESS: (a) Univ. Texas, M.D. Anderson Cancer Cent., Dep. Immunology, Box 180, 1515 Holcombe Boulevard, Hous**USA

JOURNAL: Journal of Immunology 157 (2):p700-706 1996

ISSN: 0022-1767

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: T cell coactivation is a dynamic process subject to integrin-dependent positive and negative regulation. Costimulation of human peripheral blood T cells by CD3 mAb OKT3 in conjunction with anti-alpha-4 has been shown to be down-regulated by the anti-beta-1.1 epitope-specific mAb 18D3. As expected, maximal costimulation induced by alpha-4-specific mAb L25 was inhibited (70%) by the addition of soluble mAb 18D3. Surprisingly, soluble mAb 18D3 inhibited maximal proliferation induced by the costimulatory alpha-4-beta-7-specific mAb ACT-1 by 40%, thus demonstrating that one integrin subfamily can regulate the activity of another. To determine whether mAb 18D3 could regulate more than alpha-4-associated integrin-mediated costimulation, non-alpha-4 integrins were tested. mAb 18D3 inhibited maximal proliferation induced by alpha-L-specific mAb 3D6, and an alpha-5-specific mAb 16. This clearly demonstrates that a variety of integrin costimulatory molecules (of the beta-1, beta-2, and beta-7 subfamilies) can be regulated negatively by mAb 18D3. To analyze the specificity of this negative regulation, other cell surface costimulatory molecules were tested for susceptibility to mAb 18D3. Although Abs specific for CD4, CD26, CD28, CD44, CD45RA, or CD45RO were sufficient to activate T cells when co-immobilized with anti-CD3 mAb, all were refractory to the inhibitory effects of mAb 18D3. Inhibition of T cell activation directly correlated with diminished IL-2 production. This suggests that mAb 18D3 selectively regulates integrin-dependent T cell activation by delivering a negative effect at some common point utilized by various integrin subfamilies.

4/7/5 (Item 5 from file: 5)
DIALOG(R)File 5:Biosis Previews(R)
(c) 2001 BIOSIS. All rts. reserv.

09066617 BIOSIS NO.: 199497074987

Differential expression in rheumatoid synovium and synovial fluid of alpha-4-beta-7 integrin. A novel receptor for Fibronectin and vascular cell adhesion molecule-1.

AUTHOR: Lazarovits Andrew I(a); Karsh Jacob

AUTHOR ADDRESS: (a) Univ. Hosp., Room 4TU46, Box 5339, 339 Windermere Rd., London, ON N6A 5A5**Canada

JOURNAL: Journal of Immunology 151 (11):p6482-6489 1993

ISSN: 0022-1767

DOCUMENT TYPE: Article RECORD TYPE: Abstract LANGUAGE: English

ABSTRACT: T lymphocyte adhesion to vascular endothelium plays an important role in the immunopathogenesis of rheumatoid arthritis. The migration of T lymphocytes into the synovium is mediated by a variety of adhesion molecules, notably the integrins. We have prepared Act 1, a murine mAb that identifies a novel integrin termed alpha-4-beta-7. The natural ligands for alpha-4-beta-7 are vascular cell adhesion molecule-1 and fibronectin; both molecules are upregulated in the rheumatoid synovium. We investigated the expression of alpha-4-beta-7 in the three compartments of rheumatoid arthritis, the peripheral blood, synovial fluid, and synovial membrane, utilizing the FACS and immunoperoxidase microscopy of frozen tissues. The results of our experiments show a striking differential expression of alpha-4-beta-7 integrin in rheumatoid arthritis. Sixty-two percent of synovial membrane T cells expressed high density alpha-4-beta-7, in contrast to only 4.7% of synovial fluid and 9.1% of PBL. These data suggest that the expression of alpha-4-beta-7 integrin may provide a mechanism whereby certain T cells adhere to rheumatoid synovium while others remain in the synovial fluid. The augmented expression of alpha-4-beta-7 in the synovial membrane T cells may contribute to the development and perpetuation of rheumatoid arthritis.

4/7/6 (Item 6 from file: 5)
DIALOG(R)File 5:Biosis Previews(R)
(c) 2001 BIOSIS. All rts. reserv.

08920904 BIOSIS NO.: 199396072405

CD53, a protein with four membrane-spanning domains, mediates signal transduction in human monocytes and B cells.

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ABSTRACT: CD53 is a member of a novel family of molecules with four presumably membrane-spanning domains. The structure and functional characteristics of these molecules indicate that they may play an important role in transmembrane communication. We therefore investigated whether CD53 is involved in activation of human leukocytes. Cross-linking of cell-bound F(ab')-2 fragments of two different anti-CD53 mAb with F(ab')-2 anti-mouse Iq led to cytoplasmic calcium fluxes in B cells, monocytes, and granulocytes and activation of the monocyte oxidative burst. These responses were specific for CD53, as cross-linking of CD11a, CD18, CD35, CD43, CD44, CD45, or CDw50 did not induce leukocyte activation. Low concentrations of staurosporine (10 to 20 nM) completely inhibited PMA-mediated activation, but had no effect on CD53-mediated calcium fluxes and inhibited only partially CD53-mediated oxidative burst. This suggests that CD53-mediated signaling is largely independent of protein kinase C. CD53-mediated calcium fluxes were inhibited by high concentrations of staurosporine (300 to 500 nM) but not by ADP-ribosylating toxins, suggesting dependence on tyrosine kinases rather than GTP-binding proteins. The results indicate that CD53, like several other leukocyte Ag with four membrane-spanning regions, has the ability to mediate cell activation, and support the view that these molecules are involved in transmembrane communication.

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Selective expression of integrin alpha-4-beta-7 on a subset of human CD4 positive memory T cells with hallmarks of gut-trophism.

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ABSTRACT: Human memory CD4+ T lymphocytes are heterogenous in expression of integrins; one subset has the unexpected phenotype beta-1-low-alpha-4-high. We demonstrate that this subset is unique among CD4+ cells in expression of high levels of alpha-4-beta-7, detected by a distinctive mAb Act-1. alpha-4-beta-7 is involved in binding to both fibronectin and vascular cell adhesion molecule-1; Act-1 blocks cell binding to the former and augments binding to the latter. Act-1 expression marks a subset of memory cells that, unlike the predominant circulating memory cell, has up-regulated beta-7 rather than beta-1. Their phenotype is distinct from that described for skin-homing T cells and is fully consistent with that described for gut-homing T cells. Differential adhesion capacity of this subset is verified by selective binding to FN and vascular cell adhesion molecule-1 in a beta-1-independent fashion. Thus, alpha-4-beta-7 detected on this subset of circulating normal T cells fits the expectations for a gut-homing receptor.

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Expression and function of alphainf 4/betainf 7 integrin on human natural killer cells

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In this report, we have analysed the expression and function of the alphainf 4/betainf 7 heterodimer in human natural killer (NK) cells. The expression of alphainf 4betainf 7 is induced in NK cells upon activation, as the anti-alphainf 4betainf 7 ACT-1 monoclonal antibody (mAb) faintly stained a minority of peripheral blood NK cells, whereas it strongly reacted with in viti o long-term interleukin-2 (IL-2)-activated NK cells. Incubation with ACT-1 or its F(ab')inf 2 fragments induced a strong homotypic adhesion of NK cells, comparable to that stimulated by the anti-alphainf 4 (HP1/7) mAb. Cell-cell interaction induced by the ACT-1 mAb was only prevented by another anti-alphainf 4 mAb (HP2/1) that recognizes a different epitope. In alphainf 4betainf 7-mediated cell aggregation, the alphainf 4betainf 7 heterodimer was redistributed to intercellular contact sites, thus, suggesting a direct involvement of this interprin in the formation of cell clusters. In NK cells attached to Fibronectin (FN38) or vascular cell

adhesion molecule-1 (VCAM-1), both alphainf 4betainf 7 and alphainf 4/betainf 1 integrins were redistributed at the ventral cellular membrane forming discrete contact sites. The ACT-1 mAb only partially blocked NK cell binding to FN38, but in combination with the anti-betainf 1 mAb LIA1/2, NK cell binding to FN38 was completely inhibited. In contrast, ACT-1 did not modify NK cell adhesion to VCAM-1, thus supporting the theory that the alphainf 4betainf 7 binding sites for both ligands appear to be different. Our results indicate that upon IL-2-activation, expression of functional alphainf 4/betainf 7 integrin is induced on NK cells, potentially participating in their interaction with both extracellular matrix and endothelial cells.

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 ${\bf Integrin}$ alpha 4 beta 7 co-stimulation of human peripheral blood ${\bf T}$ cell proliferation.

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The integrin alpha 4 beta 7 mediates lymphocytes adhesion to VCAM-1 on activated endothelium, fibronectin in the extracellular matrix, and the mucosal vascular addressin MAdCAM-1. It is unclear whether alpha 4 beta 7 performs any function beyond directing specific adhesion reactions. We addressed the possibility that triggering of alpha 4 beta 7 with a specific monoclonal antibody was capable of delivering an accessory stimulus that would coactivate T cells and lead to proliferation. At submitogenic levels of anti-CD3 stimulation, triggering of alpha 4 beta 7 by immobilized mAb ACT-1 resulted in T cell blastogenesis, IL-2 production, expression of the IL-2 receptor alpha chain CD25, and ultimately DNA synthesis. These results indicate that the integrin alpha 4 beta 7 is involved in more than lymphocyte adhesion and homing but also plays a role in cell signaling.

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Human cytotrophoblastic antigens defined by monoclonal antibodies.

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Tohoku journal of experimental medicine (JAPAN) Mar 1988, 154 (3) p245-51, ISSN 0040-8727 .Journal Code: VTF

Languages: ENGLISH

Document type: JOURNAL ARTICLE

Monoclonal antibodies have been raised against cytotrophoblast. Two antigens, different defined on cytotrophoblast but syncytiotrophoblast were designated ACT-1 and ACT-2, respectively. Chorionic villi were taken from normal early pregnancy and processed for immunization by two different procedures. ACT-1 was demonstrated to be present in lung alveolar cells, endothelial mucosa of the jejunum, colon, ureter, urinary bladder and the fallopian tube, and endometrial gland of the pregnant uterus. On the other hand, ACT-2 was present in the endothelial mucosa of the stomach, endothelium of the renal

vessel, and the decidua of the pregnant uterus. Although the monoclonal antibodies did not react with such established cell lines as Bewo, SCH, OVK-18, HHUA, MK-01, FL, BHK and P3 X 63Ag 8 . 653, they did react with some of the cell lines when the cell membrane was destroyed with Triton membrane but in the cytoplasm. The antigens might be shed or may disappear in the process of differentiation from cytotrophoblast to

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